

THAT WHICH IS CLAIMED:

1. A polymer composition, comprising a mixture of a polymer derivative having the structure R-O-POLY-R' and a polymer derivative having the structure R-O-POLY-O-  
5 R, wherein POLY is a water-soluble and non-peptidic polymer, R is an alkyl or an aryl group, and R' is a functional group.

2. The polymer composition of Claim 1, wherein POLY is selected from poly(alkylene oxides), poly(acryloylmorpholine), poly(oxazoline), and  
10 poly(vinylpyrrolidone).

3. The polymer composition of Claim 1, wherein POLY is poly(ethylene glycol).

4. The polymer composition of Claim 3, wherein POLY has the formula  
15  $-\text{CH}_2\text{CH}_2-\text{O}-(\text{CH}_2\text{CH}_2\text{O})_n-\text{CH}_2\text{CH}_2-$ , where n is from about 8 to about 4000.

5. The polymer composition of Claim 1, wherein R is methyl.

6. The polymer composition of Claim 1, wherein R' is selected from the group  
20 consisting of hydroxyl, mesylate, tosylate, tresylate,  $-\text{O}-\text{CO}_2\text{R}_3$  wherein  $\text{R}_3$  is H, alkyl or N-succinimidyl,  $-\text{O}-(\text{CH}_2)_n-\text{CO}_2\text{R}_3$  wherein n is 1-6 and  $\text{R}_3$  is H, alkyl or N-succinimidyl,  $-\text{NHR}_4$  wherein  $\text{R}_4$  is H or alkyl or an amine protecting group,  $-\text{O}-(\text{CH}_2)_n-\text{CH}(\text{ZR}_5)_2$  wherein n is 1-6, and Z is O or S,  $\text{R}_5$  is H or an alkyl group,  $\text{Ar}-\text{CH}=\text{CH}-\text{CH}=\text{CH}-\text{CO}_2-$ , wherein Ar represents a moiety selected from the group consisting of phenyl, substituted phenyl, biphenyl, substituted biphenyl, polycyclic aryls, substituted polycyclic aryls, and  
25 heterocyclic aryls,  $-\text{O}-(\text{CH}_2)_n-\text{CHO}$  wherein n is 1-6,  $-\text{O}_2\text{CCH}_2\text{CH}_2\text{CO}_2\text{R}_6$ , wherein  $\text{R}_6$  is H or N-succinimidyl,  $\text{CH}_2=\text{CH}-\text{CO}_2-$ , and  $-\text{O}-\text{CH}_2-\text{CO}_2\text{CH}(\text{CH}_3)\text{CH}_2\text{CO}_2-\text{NHS}$ , wherein NHS is N-succinimidyl.

7. The polymer composition of Claim 1, wherein R' is hydroxyl.  
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8. The polymer composition of Claim 1, wherein R' is -O-CO<sub>2</sub>R<sub>3</sub> or -O-(CH<sub>2</sub>)<sub>n</sub>-CO<sub>2</sub>R<sub>3</sub>, wherein n is 1-6 and R<sub>3</sub> is H, alkyl or N-succinimidyl.

9. The polymer composition of Claim 8, wherein R<sub>3</sub> is N-succinimidyl.

10. The polymer composition of Claim 1, wherein R' is -O-(CH<sub>2</sub>)<sub>n</sub>-CHO wherein n is 1-6.

11. The polymer composition of Claim 1, wherein R' is -O-(CH<sub>2</sub>)<sub>n</sub>-CH(ZR<sub>5</sub>)<sub>2</sub> wherein n is 1-6, Z is O or S, and R<sub>5</sub> is H or an alkyl group.

12. The polymer composition of Claim 1, wherein POLY is poly(ethylene glycol), R is methyl, and R' is -O-(CH<sub>2</sub>)<sub>n</sub>-CHO wherein n is 2.

13. The polymer composition of Claim 12, wherein POLY has the formula -CH<sub>2</sub>CH<sub>2</sub>-O-(CH<sub>2</sub>CH<sub>2</sub>O)<sub>n</sub>-CH<sub>2</sub>CH<sub>2</sub>-, where n is from about 8 to about 4000.

14. The polymer composition of Claim 1, wherein POLY is poly(ethylene glycol), R is methyl, and R' is hydroxyl.

15. The polymer composition of Claim 14, wherein POLY has the formula -CH<sub>2</sub>CH<sub>2</sub>-O-(CH<sub>2</sub>CH<sub>2</sub>O)<sub>n</sub>-CH<sub>2</sub>CH<sub>2</sub>-, where n is from about 8 to about 4000.

16. A polymer composition, comprising a polymer derivative having the structure R-O-POLY-R', wherein POLY is a water-soluble and non-peptidic polymer, R is an alkyl or an aryl group, and R' is a functional group, in the absence of HO-POLY-OH.

17. The polymer composition of Claim 16, wherein POLY is selected from poly(alkylene oxides), poly(acryloylmorpholine), poly(oxazoline), and poly(vinylpyrrolidone).

18. The polymer composition of Claim 16, wherein POLY is poly(ethylene glycol).

19. The polymer composition of Claim 18, wherein POLY has the formula  
5 -CH<sub>2</sub>CH<sub>2</sub>-O-(CH<sub>2</sub>CH<sub>2</sub>O)<sub>n</sub>-CH<sub>2</sub>CH<sub>2</sub>-, where n is from about 8 to about 4000.

20. The polymer composition of Claim 16, wherein R is methyl.

21. The polymer composition of Claim 16, wherein R' is selected from the group  
10 consisting of hydroxyl, mesylate, tosylate, tresylate, -O-CO<sub>2</sub>R<sub>3</sub> wherein R<sub>3</sub> is H, alkyl or N-succinimidyl, -O-(CH<sub>2</sub>)<sub>n</sub>-CO<sub>2</sub>R<sub>3</sub> wherein n is 1-6 and R<sub>3</sub> is H, alkyl or N-succinimidyl, -NHR<sub>4</sub> wherein R<sub>4</sub> is H or alkyl or an amine protecting group, -O-(CH<sub>2</sub>)<sub>n</sub>-CH(ZR<sub>5</sub>)<sub>2</sub> wherein n is 1-6, and Z is O or S, R<sub>5</sub> is H or an alkyl group, Ar-CH=CH-CH=CH-CO<sub>2</sub>-, wherein Ar represents a moiety selected from the group consisting of phenyl, substituted phenyl, biphenyl, substituted biphenyl, polycyclic aryls, substituted polycyclic aryls, and heterocyclic aryls, -O-(CH<sub>2</sub>)<sub>n</sub>-CHO wherein n is 1-6, -O<sub>2</sub>CCH<sub>2</sub>CH<sub>2</sub>CO<sub>2</sub>R<sub>6</sub>, wherein R<sub>6</sub> is H or N-succinimidyl, CH<sub>2</sub>=CH-CO<sub>2</sub>-, and -O-CH<sub>2</sub>-CO<sub>2</sub>CH(CH<sub>3</sub>)CH<sub>2</sub>CO<sub>2</sub>-NHS, wherein NHS is N-succinimidyl.  
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22. The polymer composition of Claim 16, wherein R' is hydroxyl.  
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23. The polymer composition of Claim 16, wherein R' is -O-CO<sub>2</sub>R<sub>3</sub> or  
-O-(CH<sub>2</sub>)<sub>n</sub>-CO<sub>2</sub>R<sub>3</sub>, wherein n is 1-6 and R<sub>3</sub> is H, alkyl or N-succinimidyl.

24. The polymer composition of Claim 23, wherein R<sub>3</sub> is N-succinimidyl.  
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25. The polymer composition of Claim 16, wherein R' is -O-(CH<sub>2</sub>)<sub>n</sub>-CHO  
wherein n is 1-6.

26. The polymer composition of Claim 16, wherein R' is -O-(CH<sub>2</sub>)<sub>n</sub>-CH(ZR<sub>5</sub>)<sub>2</sub>  
wherein n is a number of 1-6, Z is O or S, and R<sub>5</sub> is H or an alkyl group.  
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27. The polymer composition of Claim 16, wherein POLY is poly(ethylene glycol), R is methyl, and R' is  $-O-(CH_2)_n-CHO$ , wherein n is 2.

5 28. The polymer composition of Claim 27, wherein POLY has the formula  $-CH_2CH_2-O-(CH_2CH_2O)_n-CH_2CH_2-$ , where n is from about 8 to about 4000.

29. The polymer composition of Claim 16, wherein POLY is poly(ethylene glycol), R is methyl, and R' is hydroxyl.

10 30. The polymer composition of Claim 29, wherein POLY has the formula  $-CH_2CH_2-O-(CH_2CH_2O)_n-CH_2CH_2-$ , where n is from about 8 to about 4000.

15 31. A polymer conjugate composition, comprising a mixture of a conjugated polymer having the structure  $R-O-POLY-M_1$ , wherein POLY is a water-soluble and non-peptidic polymer, R is an alkyl or an aryl group, and  $M_1$  is a macromolecule, with a polymer derivative having the structure  $R-O-POLY-O-R$ , wherein R and POLY are as defined above.

20 32. The polymer conjugate composition of Claim 31, wherein  $M_1$  is selected from the group consisting of proteins, peptides, lipids, drugs, and polysaccharides.

25 33. The polymer conjugate composition of Claim 31, wherein POLY is selected from poly(alkylene oxides), poly(acryloylmorpholine), poly(oxazoline), and poly(vinylpyrrolidone).

34. The polymer conjugate composition of Claim 31, wherein POLY is poly(ethylene glycol).

30 35. The polymer conjugate composition of Claim 34, wherein POLY has the formula  $-CH_2CH_2-O-(CH_2CH_2O)_n-CH_2CH_2-$ , where n is from about 8 to about 4000.

36. The polymer conjugate composition of Claim 31, wherein R is methyl.

37. A method of forming a polymer conjugate composition, comprising:

5 providing a mixture of a polymer derivative having the structure R-O-POLY-R', wherein POLY is a water-soluble and non-peptidic polymer, R is an alkyl or an aryl group, and R' is a functional group, with a polymer derivative having the structure R-O-POLY-O-R, wherein R and POLY are as defined above; and

10 reacting the functional group R' with a macromolecule to form a mixture of a conjugated polymer having the formula R-O-POLY-M<sub>1</sub>, wherein POLY and R are as defined above and M<sub>1</sub> is a macromolecule, and a polymer derivative having the structure R-O-POLY-O-R.

15 38. The method of Claim 37, wherein POLY is selected from poly(alkylene oxides), poly(acryloylmorpholine), poly(oxazoline), and poly(vinylpyrrolidone).

39. The method of Claim 37, wherein POLY is poly(ethylene glycol).

20 40. The method of Claim 39, wherein POLY has the formula -CH<sub>2</sub>CH<sub>2</sub>-O-(CH<sub>2</sub>CH<sub>2</sub>O)<sub>n</sub>-CH<sub>2</sub>CH<sub>2</sub>-, where n is from about 8 to about 4000.

41. The method of Claim 37, wherein R is methyl.

25 42. The method of Claim 37, wherein R' is selected from the group consisting of hydroxyl, mesylate, tosylate, tresylate, -O-CO<sub>2</sub>R<sub>3</sub> wherein R<sub>3</sub> is H, alkyl or N-succinimidyl, -O-(CH<sub>2</sub>)<sub>n</sub>-CO<sub>2</sub>R<sub>3</sub> wherein n is 1-6 and R<sub>3</sub> is H, alkyl or N-succinimidyl, -NHR<sub>4</sub> wherein R<sub>4</sub> is H or alkyl or an amine protecting group, -O-(CH<sub>2</sub>)<sub>n</sub>-CH(ZR<sub>5</sub>)<sub>2</sub> wherein n is 1-6, and Z is O or S, R<sub>5</sub> is H or an alkyl group, Ar-CH=CH-CH=CH-CO<sub>2</sub>-, wherein Ar represents a moiety selected from the group consisting of phenyl, substituted phenyl, biphenyl, substituted biphenyl, polycyclic aryls, substituted polycyclic aryls, and  
30 heterocyclic aryls, -O-(CH<sub>2</sub>)<sub>n</sub>-CHO wherein n is 1-6, -O<sub>2</sub>CCH<sub>2</sub>CH<sub>2</sub>CO<sub>2</sub>R<sub>6</sub>, wherein R<sub>6</sub> is

H or N-succinimidyl,  $\text{CH}_2=\text{CH}-\text{CO}_2^-$ , and  $-\text{O}-\text{CH}_2-\text{CO}_2\text{CH}(\text{CH}_3)\text{CH}_2\text{CO}_2-\text{NHS}$ , wherein NHS is N-succinimidyl.

43. The method of Claim 37, wherein R' is hydroxyl.

44. The method of Claim 37, wherein R' is  $-\text{O}-\text{CO}_2\text{R}_3$  or  $-\text{O}-(\text{CH}_2)_n-\text{CO}_2\text{R}_3$ , wherein n is 1-6 and R<sub>3</sub> is H, alkyl or N-succinimidyl.

45. The method of Claim 44, wherein R<sub>3</sub> is N-succinimidyl.

46. The method of Claim 37, wherein R' is  $-\text{O}-(\text{CH}_2)_n-\text{CHO}$  wherein n is 1-6.

47. The method of Claim 37, wherein R' is  $-\text{O}-(\text{CH}_2)_n-\text{CH}(\text{ZR}_5)_2$  wherein n is a number of 1-6, Z is O or S, and R<sub>5</sub> is H or an alkyl group.

48. The method of Claim 37, wherein POLY is poly(ethylene glycol), R is methyl, and R' is  $-\text{O}-(\text{CH}_2)_n-\text{CHO}$  wherein n is 2.

49. The method of Claim 48, wherein POLY has the formula  $-\text{CH}_2\text{CH}_2-\text{O}-(\text{CH}_2\text{CH}_2\text{O})_n-\text{CH}_2\text{CH}_2-$ , where n is from about 8 to about 4000.

50. The method of Claim 37, wherein the macromolecule is selected from the group consisting of proteins, peptides, lipids, drugs, and polysaccharides.